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## INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

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| <b>(51) International Patent Classification <sup>6</sup> :</b><br>A61K 31/445, 31/395, 31/44, 31/425, 31/40,<br>31/185, 31/20, 31/60, 31/62, 31/56, 31/16, 31/19   |  | <b>A3</b> | <b>(11) International Publication Number:</b> <b>WO 98/19649</b>   |
|  |  |           | <b>(43) International Publication Date:</b> 14 May 1998 (14.05.98)   |
| <b>(21) International Application Number:</b> PCT/US97/20116<br><b>(22) International Filing Date:</b> 4 November 1997 (04.11.97)<br><br><b>(30) Priority Data:</b><br>60/028,708 5 November 1996 (05.11.96) US<br>08,963,058 3 November 1997 (03.11.97) US<br><br><b>(71) Applicant:</b> THE CHILDREN'S MEDICAL CENTER CORPORATION [US/US]; 300 Longwood Avenue, Boston, MA 02115 (US).<br><br><b>(72) Inventor:</b> D'AMATO, Robert, J.; 65 Huron Avenue, Cambridge, MA 02138 (US).<br><br><b>(74) Agents:</b> MERCHANT, Mary, Anthony et al.; Jones & Askew, LLP, 37th floor, 191 Peachtree Street, N.E., Atlanta, GA 30303 (US). |  |           | <b>(81) Designated States:</b> AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, ARIPO patent (GH, KE, LS, MW, SD, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG).<br><br><b>Published</b><br><i>With international search report.</i><br><i>Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.</i><br><br><b>(88) Date of publication of the international search report:</b><br>25 June 1998 (25.06.98) |
| <b>(54) Title:</b> METHODS AND COMPOSITIONS FOR INHIBITION OF ANGIOGENESIS   |  |           |  |
| <b>(57) Abstract</b><br><br>The present invention comprises a group of compounds that effectively inhibit angiogenesis. More specifically, thalidomide and various related compounds such as thalidomide precursors, analogs, metabolites and hydrolysis products have been shown to inhibit angiogenesis and to treat disease states resulting from angiogenesis. Additionally, antiinflammatory drugs, such as steroids and NSAIDs can inhibit angiogenesis dependent diseases either alone or in combination with thalidomide and related compounds. Importantly, these compounds can be administered orally.                     |  |           |  |

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## INTERNATIONAL SEARCH REPORT

International application No.  
PCT/US97/20116

**A. CLASSIFICATION OF SUBJECT MATTER**

IPC(6) : Please See Extra Sheet.

US CL : Please See Extra Sheet.

According to International Patent Classification (IPC) or to both national classification and IPC

**B. FIELDS SEARCHED**

Minimum documentation searched (classification system followed by classification symbols)

U.S. : 514/323, 210, 338, 373, 411, 414, 416, 417, 553, 559, 165, 161, 179, 629, 569

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

CAPLUS, BIOSIS, MEDLINE, EMBASE- COMPOUNDS HEREIN FOR THE TREATMENT OF ANGIOGENESIS RELATED DISORDERS HEREIN

**C. DOCUMENTS CONSIDERED TO BE RELEVANT**

| Category*      | Citation of document, with indication, where appropriate, of the relevant passages                  | Relevant to claim No.               |
|----------------|---|-------------------------------------|
| X<br>----<br>Y | WO 95/04533 A2 (ANDRULIS PHARMACEUTICALS CORPORATION) 16 February 1995, entire document.            | 1-8, 10-11, 13-20<br>-----<br>9, 12 |
| Y              | WO 94/20085 A1 (CHILDREN'S HOSPITAL MEDICAL CENTER CORPORATION) 15 September 1994, entire document. | 1-20                                |

☐ Further documents are listed in the continuation of Box C. ☐ See patent family annex.

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Date of the actual completion of the international search

24 MARCH 1998

Date of mailing of the international search report

27 APR 1998

Name and mailing address of the ISA/US  
Commissioner of Patents and Trademarks  
Box PCT  
Washington, D.C. 20231

Facsimile No. (703) 305-3230

Authorized officer

M. MOEZIE

Telephone No. (703) 308-1235

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